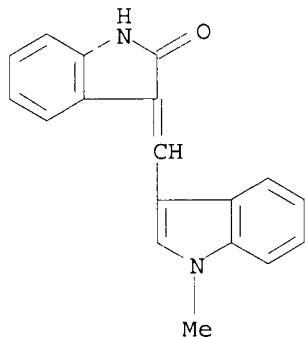


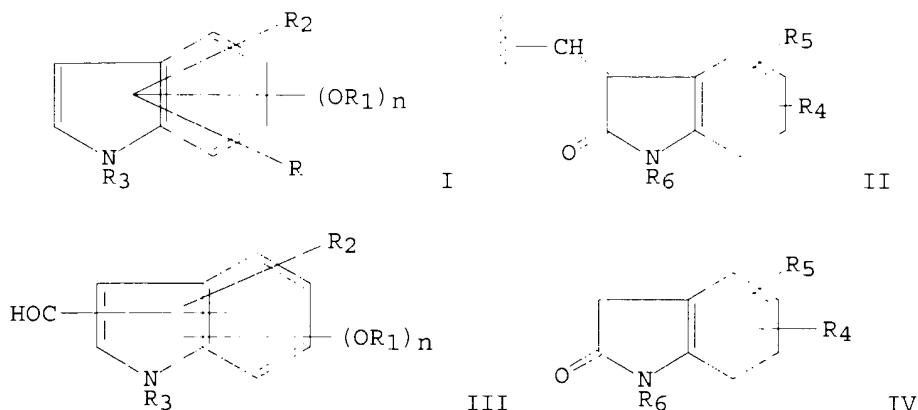
L7 ANSWER 8 OF 9 CAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1993:449225 CAPLUS  
 DOCUMENT NUMBER: 119:49225  
 TITLE: Methylenoindole derivatives and process for their preparation  
 INVENTOR(S): Buzzetti, Franco; Longo, Antonio; Colombo, Maristella  
 PATENT ASSIGNEE(S): Farmitalia Carlo Erba Srl, Italy  
 SOURCE: PCT Int. Appl., 33 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9301182	A1	19930121	WO 1992-EP1569	19920710
W: AU, CA, FI, HU, JP, KR, RU, US RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LU, MC, NL, SE				
IL 102383	A1	19970930	IL 1992-102383	19920702
CA 2091058	AA	19930113	CA 1992-2091058	19920710
EP 525472	A2	19930203	EP 1992-111757	19920710
EP 525472	A3	19930224		
R: PT				
AU 9222777	A1	19930211	AU 1992-22777	19920710
AU 656015	B2	19950119		
ZA 9205169	A	19930428	ZA 1992-5169	19920710
EP 552329	A1	19930728	EP 1992-914619	19920710
EP 552329	B1	20011004		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, NL, SE				
JP 06501494	T2	19940217	JP 1993-501981	19920710
JP 3188701	B2	20010716		
HU 67496	A2	19950428	HU 1993-723	19920710
RU 2072989	C1	19970210	RU 1993-4893	19920710
AT 206420	E	20011015	AT 1992-914619	19920710
ES 2165357	T3	20020316	ES 1992-914619	19920710
US 5409949	A	19950425	US 1993-987280	19930312
PRIORITY APPLN. INFO.:			GB 1991-15160	A 19910712
			WO 1992-EP1569	A 19920710

OTHER SOURCE(S): MARPAT 119:49225  
 GI

L7 ANSWER 9 OF 9 CAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1969:403203 CAPLUS  
 DOCUMENT NUMBER: 71:3203  
 TITLE: Indole chemistry. VI. .alpha.,.beta.'-Diindolylmethanes and .alpha.,.beta.'-diindolylmethenes  
 AUTHOR(S): Von Dobeneck, Henning; Wolkenstein, Dieter;  
 Blankenstein, Guenter  
 CORPORATE SOURCE: Tech. Hochsch. Muenchen, Munich, Fed. Rep. Ger.  
 SOURCE: Chemische Berichte (1969), 102(4), 1347-56  
 CODEN: CHBEAM; ISSN: 0009-2940  
 DOCUMENT TYPE: Journal  
 LANGUAGE: German  
 GI For diagram(s), see printed CA Issue.  
 AB Urorosein was prep'd. by polycondensation of indol-3-ylglycolic acid, followed by oxidn. with the formation of the .alpha.,.beta.'-diindolylmethene chromophore. .alpha.,.beta.'-Diindolylmethanes (I) were prep'd. by the reaction of .alpha.- and .beta.-unsubstituted indoles with glyoxylic acid; .alpha.,.beta.'-diindolylmethenes, from .alpha.-formylindoles and .beta.-unsubstituted indoles. Oxo-.beta.,.beta.'- and oxo-.alpha.,.beta.'-diindolylmethenes were prep'd. from .beta.-and .alpha.-formylindoles and oxindoles.  
 IT 22813-84-9P 22813-88-3P  
 RL: SPN (Synthetic preparation); PREP (Preparation)  
 (prepn. of)  
 RN 22813-84-9 CAPLUS  
 CN 2H-Indol-2-one, 1,3-dihydro-3-[(1-methyl-1H-indol-3-yl)methylene] - (9CI)  
 (CA INDEX NAME)





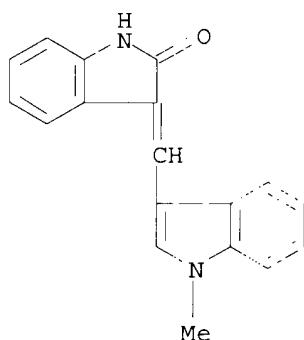
AB The title compds. I [R = II; R4 = H, OH, C1-6 alkoxy, C2-6 alkanoyloxy, CO2H, NO2, NHR7 (R7 = H, C1-6 alkyl); R5 = H, C1-6 alkyl or halo; R6 = H, C1-6 alkyl; n = 0-2; R1 = H, C1-6 alkyl, C2-6 alkanoyl; R2 = H, C1-6 alkyl, halo, CN, CO2H, NO2, NHR7; R3 = H, C1-6 alkyl, C2-6 alkanoyl] were prep'd. by condensation of aldehydes III with oxindoles IV, and were evaluated as tyrosine kinase inhibitors. Thus, a soln. of 145 mg 3-indolecarboxaldehyde, 149 mg 5-hydroxy-2-oxindole, and 60 mg piperidine in 10 mL abs. EtOH was heated for 3 h at 60.degree.; workup afforded 60% I (R in 3-position, n = 0, R2 = R3 = R5 = R6 = H, R4 = 5-OH) (V). IC50 (.mu.M) values for V of 0.4 were detd. for both the myelin basic protein phosphorylation assay and the autophosphorylation assay. A 0.150 g tablet formulation contg. 25 mg active substance and lactose, corn starch, talc powder, and magnesium stearate is given.

IT 22813-84-9P

RL: SPN (Synthetic preparation); PREP (Preparation)  
(prepn. of, as tyrosine kinase inhibitor)

RN 22813-84-9 CAPLUS

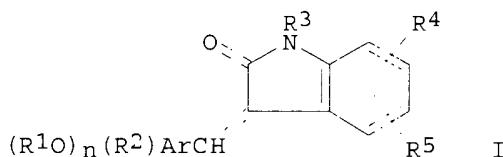
CN 2H-Indol-2-one, 1,3-dihydro-3-[(1-methyl-1H-indol-3-yl)methylene]- (9CI)  
(CA INDEX NAME)



L7 ANSWER 7 OF 9 CAPLUS COPYRIGHT 2003 ACS  
 ACCESSION NUMBER: 1995:813058 CAPLUS  
 DOCUMENT NUMBER: 123:208831  
 TITLE: Biologically active 3-substituted oxindole derivatives useful as anti-angiogenic agents  
 INVENTOR(S): Heath, William Francis Heat, Jr.; McDonald, John Hampton III; Brasca, Maria Gabriella; Orzi, Fabrizio; Crugnola, Angelo; Ballinari, Dario; Mariani, Mariangela  
 PATENT ASSIGNEE(S): Pharmacia S.P.A., Italy  
 SOURCE: PCT Int. Appl., 104 pp.  
 CODEN: PIXXD2  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 9517181	A1	19950629	WO 1994-EP3664	19941108
W: AU, BY, CA, HU, JP, KR, KZ, NO, PL, RU, UA RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
CA 2155098	AA	19950629	CA 1994-2155098	19941108
AU 9480612	A1	19950710	AU 1994-80612	19941108
AU 676958	B2	19970327		
EP 684820	A1	19951206	EP 1994-931583	19941108
EP 684820	B1	20010816		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, NL, PT, SE				
HU 73176	A2	19960628	HU 1995-2761	19941108
JP 08507089	T2	19960730	JP 1994-517121	19941108
AT 204168	E	20010915	AT 1994-931583	19941108
ES 2162871	T3	20020116	ES 1994-931583	19941108
ZA 9410204	A	19951110	ZA 1994-10204	19941212
US 5576330	A	19961119	US 1994-354215	19941212
IL 112010	A1	19981030	IL 1994-112010	19941216
NO 9503146	A	19950810	NO 1995-3146	19950810
PRIORITY APPLN. INFO.:			GB 1993-26136	A 19931222
			WO 1994-EP3664	W 19941108

OTHER SOURCE(S): MARPAT 123:208831  
 GI



AB Compds. I (Ar = naphthalene, tetralin, quinoline, isoquinoline, indole; n = 0 or an integer of 1 to 3; R<sup>1</sup> = H, C<sub>1-6</sub> alkyl, C<sub>2-6</sub> alkanoyl; R<sup>2</sup> = H, halogen, C<sub>1-6</sub> alkyl, cyano, carboxy, nitro, NHR; R = H, C<sub>1-6</sub> alkyl; R<sup>3</sup> = H, C<sub>1-6</sub> alkyl; R<sup>4</sup> = H, OH, C<sub>1-6</sub> alkoxy, C<sub>2-6</sub> alkanoyloxy, carboxy, nitro, NHR; R<sup>5</sup> = H, C<sub>1-6</sub> alkyl, halogen) or a pharmaceutically acceptable salt

thereof are useful as angiogenesis inhibitors. Products contg. an angiogenesis inhibitor or a pharmaceutically acceptable salt thereof and an antitumor agent are used as a combined prepn. for anticancer therapy. A compn. (for 10,000 tablets) contg. 3-[(3'-hydroxy-2'-tetralyl)methylene]-2-oxindole 250, lactose 800, corn starch 415, talc 30 and Mg stearate 5 g, resp., was formulated.

## IT 22813-84-9

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(oxindole derivs. as anti-angiogenic agents)

RN 22813-84-9 CAPLUS

CN 2H-Indol-2-one, 1,3-dihydro-3-[(1-methyl-1H-indol-3-yl)methylene] - (9CI)  
(CA INDEX NAME)